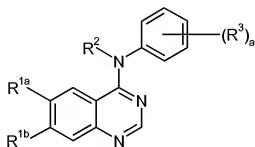


**CLAIM AMENDMENTS:**

**This listing of claims will replace all prior versions and listing of claims in the application.**

**Listing of the Claims:**

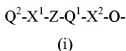
Claim 1 (**currently amended**): A quinazoline derivative of the Formula I:



I

wherein:

one of R<sup>1a</sup> or R<sup>1b</sup> is a group of sub-formula (i)



where X<sup>2</sup> and X<sup>1</sup> are independently selected from a direct bond or a group -[CR<sup>4</sup>R<sup>5</sup>]<sub>m</sub>, wherein m is an integer from 1 to 6,

Z is C(O), SO<sub>2</sub>, -C(O)NR<sup>10</sup>-, -N(R<sup>10</sup>)C(O)-, -C(O)O- or -OC(O)- where R<sup>10</sup> is hydrogen or (1-6C)alkyl,

and each of R<sup>4</sup> and R<sup>5</sup> is independently selected from hydrogen, hydroxy, (1-4C)alkyl, halo(1-4C)alkyl, hydroxy (1-4C)alkyl, (1-4C)alkoxy(1-4C)alkyl, or R<sup>4</sup> and R<sup>5</sup> together with the carbon atom(s) to which they are attached form a (3-7)cycloalkyl ring, provided that when a group R<sup>4</sup> or R<sup>5</sup> is hydroxy, m is at least 2 and the carbon atom to which the hydroxy group is attached is not also attached to another oxygen or a nitrogen atom;

Q<sup>1</sup> is a piperidinyl ring, which is optionally substituted by one or two substituents selected from halogeno, trifluoromethyl, trifluoromethoxy, cyano, nitro, hydroxy, amino, carboxy, carbamoyl,

acryloyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (2-6C)alkenylthio, (2-6C)alkynylthio, (1-6C)alkylsulfanyl, (2-6C)alkenylsulfanyl, (2-6C)alkynylsulfanyl, (1-6C)alkylsulfonyl, (2-6C)alkenylsulfonyl, (2-6C)alkynylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, sulfamoyl, N-(1-6C)alkylsulfamoyl, N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino, N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, carbamoyl(1-6C)alkyl, N-(1-6C)alkylcarbamoyl(1-6C)alkyl, N,N-di-[(1-6C)alkyl]carbamoyl(1-6C)alkyl, sulfamoyl(1-6C)alkyl, N-(1-6C)alkylsulfamoyl(1-6C)alkyl, N,N-di-[(1-6C)alkyl]sulfamoyl(1-6C)alkyl, (2-6C)alkanoyl(1-6C)alkyl, (2-6C)alkanoyloxy(1-6C)alkyl, (2-6C)alkanoylamino(1-6C)alkyl, N-(1-6C)alkyl-(2-6C)alkanoylamino(1-6C)alkyl and (1-6C)alkoxycarbonyl(1-6C)alkyl;

Q<sup>2</sup> is an isoxazolyl ring optionally substituted by one or two groups, which may be the same or different, selected from halogeno, hydroxy, nitro, amino, cyano, carbamoyl, (1-4C)alkyl, (1-4C)alkoxy, (2-4C)alkanoyl and (1-4C)alkylsulfonyl, [(1-4C)alkyl]amino, di[(1-4C)alkyl]amino, N-[(1-4C)alkyl]carbamoyl, and N,N-di[(1-4C)alkyl]carbamoyl;

and wherein any (2-4C)alkanoyl group in a substituent on Q<sup>2</sup> optionally bears one or two substituents, which may be the same or different, selected from hydroxy and (1-3C)alkyl,

and wherein any (1-4C)alkyl group in a substituent on Q<sup>2</sup> optionally bears one or two substituents, which may be the same or different, selected from hydroxy, (1-4C)alkoxy and halogeno; one of more substituents selected from halogeno, trifluoromethyl, trifluoromethoxy, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, acryloyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (2-6C)alkenylthio, (2-6C)alkynylthio, (1-6C)alkylsulfanyl, (2-6C)alkenylsulfanyl, (2-6C)alkynylsulfanyl, (1-6C)alkylsulfonyl, (2-6C)alkenylsulfonyl, (2-6C)alkynylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, sulfamoyl, N-(1-6C)alkylsulfamoyl,

N,N-di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino,  
N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, carbamoyl(1-6C)alkyl,  
N-(1-6C)alkylcarbamoyl(1-6C)alkyl, N,N-di-[(1-6C)alkyl]carbamoyl(1-6C)alkyl,  
sulfamoyl(1-6C)alkyl, N-(1-6C)alkylsulfamoyl(1-6C)alkyl,  
N,N-di-[(1-6C)alkyl]sulfamoyl(1-6C)alkyl, (2-6C)alkanoyl(1-6C)alkyl,  
(2-6C)alkanoyloxy(1-6C)alkyl, (2-6C)alkanoylamino(1-6C)alkyl,  
N-(1-6C)alkyl-(2-6C)alkanoylamino(1-6C)alkyl and (1-6C)alkoxy-carbonyl(1-6C)alkyl,  
and wherein any (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl and (2-6C)alkanoyl  
substituent on Q<sup>1</sup> or Q<sup>2</sup> optionally bears one or more substituents which may be the same or  
different selected from halogeno, hydroxy and (1-6C)alkyl and/or optionally a substituent  
selected from cyano, nitro, carboxy, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy,  
hydroxy(1-6C)alkoxy, (1-4C)alkoxy(1-6C)alkoxy, (2-6C)alkanoyl, (2-6C)alkanoyloxy and  
NR<sup>a</sup>R<sup>b</sup>, wherein R<sup>a</sup> is hydrogen or (1-4C)alkyl and R<sup>b</sup> is hydrogen or (1-4C)alkyl, and wherein  
any (1-4C)alkyl in R<sup>a</sup> or R<sup>b</sup> optionally bears one or more substituents which may be the same or  
different selected from halogeno and hydroxy and/or optionally a substituent selected from  
cyano, nitro, (2-4C)alkenyl, (2-4C)alkynyl, (1-4C)alkoxy, hydroxy(1-4C)alkoxy and  
(1-2C)alkoxy(1-4C)alkoxy;

or R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached form a 4, 5 or 6  
membered ring, which optionally bears 1 or 2 substituents, which may be the same or different,  
on an available ring carbon atom selected from halogeno, hydroxy, (1-4C)alkyl and  
(1-3C)alkylenedioxy, and may optionally bear on any available ring nitrogen a substituent  
(provided the ring is not thereby quaternised) selected from (1-4C)alkyl, (2-4C)alkanoyl and  
(1-4C)alkylsulfonyl,

and wherein any (1-4C)alkyl or (2-4C)alkanoyl group present as a substituent on the ring  
formed by R<sup>a</sup> and R<sup>b</sup> together with the nitrogen atom to which they are attached, optionally bears  
one or more substituents which may be the same or different selected from halogeno and  
hydroxy and/or optionally a substituent selected from (1-4C)alkyl and (1-4C)alkoxy;

and wherein Q<sup>1</sup> optionally bears 1 or 2 oxo (=O) or thioxo (=S) substituents;

and the other of  $R^{1a}$  or  $R^{1b}$  is a group  $R^1$  which is hydrogen, (1-6C)alkoxy and (1-4C)alkoxy(1-6C)alkoxy, and wherein any (1-6C)alkoxy group within  $R^1$  optionally bears 1, 2 or 3 substituents, which may be the same or different, selected from hydroxy, fluoro and chloro selected from hydrogen, hydroxy, (1-6C)alkoxy, (2-6C)alkenyl, (2-6C)alkynyl, or a group of the formula :



wherein  $X^3$  is a direct bond or is selected from O or S, and  $Q^4$  is (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heterocyclyl or heterocyclyl-(1-6C)alkyl;

——— and wherein adjacent carbon atoms in any (2-6C)alkylene chain within a  $R^+$  substituent are optionally separated by the insertion into the chain of a group selected from O, S,  $SO_2$ ,  $SO_2$ ,  $N(R^4)$ , CO,  $CH(OR^4)$ ,  $CON(R^4)$ ,  $N(R^4)CO$ ,  $SO_2N(R^4)$ ,  $N(R^4)SO_2$ ,  $CH=CH$  and  $C\equiv C$  wherein  $R^4$  is hydrogen or (1-6C)alkyl;

——— and wherein any  $CH_2=CH-$  or  $HC\equiv C-$  group within a  $R^+$  substituent optionally bears at the terminal  $CH_2=$  or  $HC\equiv$  position a substituent selected from halogeno, carboxy, carbamoyl, (1-6C)alkoxycarbonyl,  $N$ -(1-6C)alkylcarbamoyl,  $N,N$ -di-[(1-6C)alkyl]carbamoyl, amino (1-6C)alkyl, (1-6C)alkylamino (1-6C)alkyl and di-[(1-6C)alkyl]amino (1-6C)alkyl or from a group of the formula :



wherein  $X^4$  is a direct bond or is selected from CO and  $N(R^5)CO$ , wherein  $R^5$  is hydrogen or (1-6C)alkyl, and  $Q^5$  is heterocyclyl or heterocyclyl-(1-6C)alkyl;

——— and wherein any alkyl or alkylene group within a  $R^+$  substituent optionally bears one or more halogeno, (1-6C)alkyl, hydroxy, cyano, amino, carboxy, carbamoyl, sulfamoyl, (1-6C)alkoxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl,  $N$ -(1-6C)alkylcarbamoyl,  $N,N$ -di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino,  $N$ -(1-6C)alkyl (2-6C)alkanoylamino,  $N$ -(1-6C)alkylsulfamoyl,  $N,N$ -di-[(1-6C)alkyl]sulfamoyl, (1-6C)alkanesulfonylamino and  $N$ -(1-6C)alkyl (1-6C)alkanesulfonylamino, or from a group of the formula:



wherein  $X^5$  is a direct bond or is selected from O, S, SO, SO<sub>2</sub>, N(R<sup>6</sup>), CO, CH(OR<sup>6</sup>), CON(R<sup>6</sup>), N(R<sup>6</sup>)C(O), SO<sub>2</sub>N(R<sup>6</sup>), N(R<sup>6</sup>)SO<sub>2</sub>, C(R<sup>6</sup>)<sub>2</sub>O, C(R<sup>6</sup>)<sub>2</sub>S and C(R<sup>6</sup>)<sub>2</sub>N(R<sup>6</sup>), wherein R<sup>6</sup> is hydrogen or (1-6C)alkyl, and Q<sup>6</sup> is (3-7C)cycloalkyl, (3-7C)cycloalkyl-(1-6C)alkyl, (3-7C)cycloalkenyl, (3-7C)cycloalkenyl-(1-6C)alkyl, heterocycyl or heterocycyl-(1-6C)alkyl,

— and wherein any heterocycyl group within a substituent on R<sup>1</sup> optionally bears 1, 2 or 3 substituents, which may be the same or different, selected from halogeno, trifluoromethyl, cyano, nitro, hydroxy, amino, carboxy, carbamoyl, formyl, mercapto, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxy-carbonyl, N-(1-6C)alkyl-carbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (2-6C)alkanoylamino, N-(1-6C)alkyl-(2-6C)alkanoylamino, N-(1-6C)alkylsulfonyl, N,N-di-[(1-6C)alkyl]sulfonyl, (1-6C)alkanesulfonylamino, and N-(1-6C)alkyl-(1-6C)alkanesulfonylamino, or from a group of the formula:



wherein  $X^6$  is a direct bond or is selected from O, N(R<sup>8</sup>) and C(O), wherein R<sup>8</sup> is hydrogen or (1-6C)alkyl, and R<sup>7</sup> is halogeno (1-6C)alkyl, hydroxy (1-6C)alkyl, carboxy (1-6C)alkyl, (1-6C)alkoxy (1-6C)alkyl, cyano (1-6C)alkyl, amino (1-6C)alkyl, (1-6C)alkylamino (1-6C)alkyl, di-[(1-6C)alkyl]amino (1-6C)alkyl, (2-6C)alkanoylamino (1-6C)alkyl, (1-6C)alkoxy-carbonylamino (1-6C)alkyl, carbamoyl (1-6C)alkyl, N-(1-6C)alkyl-carbamoyl (1-6C)alkyl, N,N-di-[(1-6C)alkyl]carbamoyl (1-6C)alkyl, (2-6C)alkanoyl (1-6C)alkyl or (1-6C)alkoxy-carbonyl (1-6C)alkyl,

— and wherein any heterocycyl group within a substituent on R<sup>1</sup> optionally bears 1 or 2 exo or thioxo substituents;

R<sup>2</sup> is selected from hydrogen and (1-6C)alkyl;

each R<sup>3</sup>, which may be the same or different, is selected from halogeno, cyano, nitro,

hydroxy, amino, carboxy, carbamoyl, sulfamoyl, trifluoromethyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylsulfinyl, (1-6C)alkylsulfonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, N-(1-6C)alkylsulfamoyl, and N,N-di-[(1-6C)alkyl]sulfamoyl

a is 1, 2 or ~~3, 3, 4 or 5~~;

or a pharmaceutically acceptable salt thereof;

subject to the proviso that the compound of formula I is not

~~N-(3,4-dichlorophenyl)-7-[(4-{(3,5-dimethylisoxazol-4-yl)carbonyl}morpholin-2-yl)methoxy]-6-methoxyquinazolin-4-amine.~~

Claim 2 (**previously presented**): The quinazoline derivative according to claim 1 wherein X<sup>2</sup> is a direct bond.

Claims 3-6 (**cancelled**).

Claim 7 (**currently amended**): The quinazoline derivative according to claim 1-6 wherein R<sup>1</sup> is selected from methoxy, ethoxy, isopropoxy, cyclopropylmethoxy, 2-hydroxyethoxy, 2-fluoroethoxy, 2-methoxyethoxy, 2,2-difluoroethoxy, 2,2,2-trifluoroethoxy or 3-hydroxy-3-methylbutoxy.

Claim 8 (**currently amended**): The quinazoline derivative according to claim 7-5 wherein R<sup>1</sup> is methoxy.

Claim 9 (**previously presented**): The quinazoline derivative according to claim 1 wherein X<sup>1</sup> is suitably a direct bond or a (1-6C)alkylene group.

Claim 10 (**previously presented**): The quinazoline derivative according to claim 9 wherein X<sup>1</sup> is a direct bond or methylene or ethylene group.

Claim 11 (**previously presented**): The quinazoline derivative according to claim 1 wherein Z is selected from -C(O)-, -NR<sup>10</sup>-C(O)- (wherein R<sup>10</sup> is H or (1-6C)alkyl), and -O-C(O)-.

Claim 12 (**previously presented**): The quinazoline derivative according to claim 11, wherein Z is -C(O)-.

Claim 13 (**previously presented**): The quinazoline derivative according to claim 11, wherein Z is selected from -NH-C(O)- and -O-C(O)-.

Claims 14-15 (**cancelled**).

Claim 16 (**currently amended**): The quinazoline derivative according to claim ~~1~~<sup>14</sup>, wherein the group Q<sup>2</sup>-X<sup>1</sup>-Z- is linked to the piperidiny1 nitrogen of Q<sup>1</sup>.

Claims 17-24 (**cancelled**).

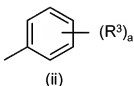
Claim 25 (**currently amended**): The quinazoline derivative according to claim ~~1~~<sup>23</sup> wherein Q<sup>2</sup> is unsubstituted or substituted by a (1-4C)alkyl group, a (1-4C)alkoxy group, halogeno, amino, nitro, cyano, carbamoyl, di-[(1-4C)alkyl]amino, and N,N-di[(1-4C)alkyl]carbamoyl.

Claim 26 (**previously presented**): The quinazoline derivative according to claim 1 wherein R<sup>2</sup> is hydrogen.

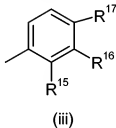
Claim 27 (**cancelled**).

Claim 28 (**previously presented**): The quinazoline derivative according to claim 1, wherein an R<sup>3</sup> is in the para position on the anilin ring, and this is selected from halogeno, cyano, nitro, hydroxy, amino, trifluoromethyl, (1-6C)alkyl, (2-8C)alkenyl, (2-8C)alkynyl, (1-6C)alkoxy, (2-6C)alkenyloxy, (2-6C)alkynyloxy, (1-6C)alkylthio, (1-6C)alkylamino and di-[(1-6C)alkyl]amino.

Claim 29 (**previously presented**): The quinazoline derivative according to claim 1 wherein the group of sub-formula (ii)



in formula (I) is a group of sub-formula (iii)



where one of R<sup>15</sup> or R<sup>17</sup> is hydrogen and the other is halogeno, and R<sup>16</sup> is halogeno.

Claim 30 (**previously presented**): The quinazoline derivative according to claim 29 wherein the group of sub-formula (iii) is 3-chloro-2-fluorophenyl, or 3-chloro-4-fluorophenyl.

Claim 31 (**currently amended**): The compound according to claim 1 selected from one of the following:

- (1) *N*-(3-chloro-2-fluorophenyl)-6- {[1-(isoxazol-5-ylcarbonyl)piperidin-4-yl]oxy}-7-methoxyquinazolin-4-amine;
- (2) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(3-methylisoxazol-5-yl)acetyl]piperidin-4-yl}oxy)quinazolin-4-amine;



- (3) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(3-methylisoxazol-5-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (4) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(5-methylisoxazol-3-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (5) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(5-methylisoxazol-4-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (6) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({1-[(3-methylisoxazol-4-yl)carbonyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (7) *N*-(3-chloro-2-fluorophenyl)-6-({1-[(3,5-dimethylisoxazol-4-yl)carbonyl]piperidin-4-yl}oxy)-7-methoxyquinazolin-4-amine;
- (8) *N*-(3-chloro-2-fluorophenyl)-7-{{1-(isoxazol-5-ylcarbonyl)piperidin-4-yl}oxy}-6-methoxyquinazolin-4-amine;
- (9) *N*-(3-chloro-2-fluorophenyl)-6-methoxy-7-({1-[(3-methylisoxazol-5-yl)acetyl]piperidin-4-yl}oxy)quinazolin-4-amine;
- (10) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-({(3*R*)-1-[(3-methylisoxazol-5-yl)acetyl]piperidin-3-yl}oxy)quinazolin-4-amine; and
- ~~(11) *N*-(3-chloro-2-fluorophenyl)-7-methoxy-6-{{{(3*R*)-1-(4-{*N,N*-dimethylcarbamoyl}-1*H*-pyrazol-1-yl)acetyl}piperidin-3-yl}oxy)}quinazolin-4-amine; and~~
- ~~(12)~~ (11) 4-({4-[(3-Chloro-2-fluorophenyl)amino]-7-methoxyquinazolin-6-yl}oxy)-*N*-(3,5-dimethylisoxazol-4-yl)piperidine-1-carboxamide.

Claims 32-33 (**cancelled**).

Claim 34 (**previously presented**): A pharmaceutical composition which comprises a quinazoline derivative of the Formula I, or a pharmaceutically-acceptable salt thereof, as defined in claim 1 in association with a pharmaceutically-acceptable diluent or carrier.

Claims 35-37 (**cancelled**).